



ATTACHMENT B
Amendments to the Specification

Please replace the paragraph at page 21, lines 6-16 with the following amended paragraph:

Seven peptides, having the following sequences, were obtained:

- 1 - X-Met-Tyr-Asp-Gly-Pro (SEQ ID No. 11)
- 2 - X-Phe-Asn-Leu-Tyr-Pro-Arg (SEQ ID No. 12)
- 3 - X-Val-Leu-Glu-Asp-Gly-Thr-Leu-His-Val-Thr-Glu-Gly (SEQ ID No. 13)
- 4 - X-Ile-Gly-X-X-Ala-Gln-Val-(His ?)-Ala-Glu-Asn-Gly-X-Ile-Ile-Ala-Glu-Glu-Gln
(SEQ ID No. 14)
- 5 - X-X-Glu-Asn-Gln-Phe-Val-Ala-Val-Thr (SEQ ID No. 15)
- 6 - X-Val-Asn-Asp-(Asp ?)-Gln-Ser-Phe-Tyr-Ala-Asp-Ile-Tyr-Met-Glu-(Asp ?)-(Gly ?)-
Leu-Ile (SEQ ID No. 16)
- 7 - X-X-X-Phe-Val-Thr-X-Pro-X-Leu-X-Pro (SEQ ID No. 17)

Please replace the paragraph at page 21, lines 26-31 with the following amended paragraph:

The cloning of the cDNA of the POP-66 protein or of related proteins was carried out by using degenerate oligonucleotide probes obtained from fragments of two peptides:

Ile-Ile-Ala-Glu-Glu-Gln (SEQ ID No. 18)

Tyr-Ala-Asp-Ile-Tyr-Met-Glu-(Asp ?) (SEQ ID No. 19)

Please replace the paragraph at page 21, lines 32-37 with the following amended paragraph:

Four sets of degenerate oligonucleotide primers (sense/antisense) are therefore determined (AT(C/T)ATTGC(T/A)GA(A/G)CA (SEQ ID No. 20);
TG(C/T)TC(T/C)AC(T/A)-GCAT(A/G)AT (SEQ ID No. 21);
TATGC(A/T)GA(C/T)AT(C/T)ATGGA (SEQ ID No. 22);
TCCAT(G/A)TA(G/A)CT-(T/A)GCATA (SEQ ID No. 23), and used for a PCR amplification.

Please replace the paragraph at page 21, line 38 to page 22, line 2 with the following amended paragraph:

The matrix is prepared in the form of double-stranded cDNA (Promega PROMEGA kit) from poly(A+)RNA extracted from the brain of rats 10 days old (Zivic-Miller, USA) using the ~~Fast-Track~~ FAST TRACK mRNA isolation kit (Invitrogen).

Please replace the paragraph at page 22, lines 15-19 with the following amended paragraph:

A comparative analysis of the nucleic acid sequences using the ~~Genbank~~ GENBANK and EMBL databases reveals that MFB-17 is a partial cDNA with a nucleotide sequence identical to that of a segment of TOAD-64, a rat neuronal protein (Minturn et al., 1995).

Please replace the paragraph at page 22, lines 32-37 with the following amended paragraph:

To obtain a complete TOAD-64 protein, the ds-cDNA matrix of rat brains was amplified with two sets of primers situated at the 5' and 3' extremities of the coding regions (sense: GGCATATGTCTTATCAGGGGAAG (SEQ ID No. 24); antisense: GCGAATTCTTAGCCCAGGCTGATG (SEQ ID No. 25)).

Please replace the paragraph at page 27, lines 33-35 with the following amended paragraph:

Anti-peptide 4 antibodies directed against the peptide LEDGTLHVTEGS (SEQ ID No. 26) were produced according to the same protocol.

Please replace the paragraph at page 35, lines 10-27 with the following amended paragraph:

A flag (EcoRI-ATGGACTACAAGGACGACGATGACAAGG-BamHI) (SEQ ID No. 27) sequence (Kodak) was cloned in the EcoRI site of pSG5 followed by ULIP-1 (EMBEL X87817), base pairs: 309-2023), ULIP-2 (Y10339, base pairs: 23-1741), ULIP-3 (Y09080, base pairs: 269-1991) or ULIP-4 (Y09079, base pairs: 102-1820), respectively. The HeLa cells were cultured in DMEM media (Gibco) to which 10% of foetal calf serum (v/v) was added. The transfections were carried out by calcium phosphate precipitation (Maniatis et al., 1978). The HeLa cells were mixed with 5 µg of Psg5FLAG-ulip-1, 2, 3 and 4 plasmids and 10 µg of pUC18. Twenty-four hours after the transfection, the HeLa cells were fixed with 4% paraformaldehyde and immunolabelled with different human sera (dilution 1/300), visualized by human anti-IgG antibodies

conjugated to FITC (Biosys), or anti-flag antibodies (M2, Kodak) (dilution 1/1000), visualized by anti-rabbit antibodies conjugated to Texas red (Vector).

Please replace the paragraph at page 37, lines 4-8 with the following amended paragraph:

PCD: paraneoplastic cerebellar degeneration;

LE: ~~lymbic~~limbic encephalitis;

PEM: paraneoplastic encephalomyelitis;

UC: undifferentiated carcinoma;

SCLC: small-cell lung carcinoma.

Please replace the paragraphs at page 39, lines 3-11 with the following amended paragraph:

Composition of the oligonucleotide probes used for ULIP-3 PCR

5' ATAGAGGAGCGGATGACG (899) (SEQ ID No. 28) 3'

GCTGTTATGGTCTTCAACTTGTCGG (SEQ ID No. 29) (1092)

GGCCTGTTATGGTCTTCAACTTGTCG (SEQ ID No. 30) (1093)

Composition of the oligonucleotide probes used for ULIP-2 PCR

5' AGGAGGAGTGAAGACCATCG (SEQ ID No. 31) (5227) 3'

CTTATGCCACTCGCTGATGTCC (SEQ ID No. 32) (509).

Please replace the paragraphs at page 39, line 26 to page 40, line 2 with the following amended paragraph:

The total RNAs are extracted from cerebral tumours preserved in liquid nitrogen according to the conventional RNAZOL™ technique (Bioprobe, France). Reverse

transcription was carried out using oligo(dt)₁₈ on 1 µg of total RNA and the PCR was carried out with 1/20 of the volume of the mixture for the reverse transcription (RT mix).

The primers used for ULIP-4 are: 5'CATCTGGCTGTCGCTGCAC3' (SEQ ID No. 33), 5'GCCGCCCCTACCAGAGACC3' (SEQ ID No. 34), and for GAPDH:

5'GGAGATTCAGTGTGGTGG3' (SEQ ID No. 35), 5'GGCTCTCCAGAACATCATCC3' (SEQ ID No. 36). The cDNA was denatured at 95°C for five minutes. PCR

amplification was carried out for 30 cycles. ULIP-4: 95°C, 45 sec; 62°C, 45 sec; 72°C, 45 sec. GAPDH: 95°C, 45 sec; 55°C, 45 sec; 72°C, 45 sec. The final extension was carried out at 72°C for 5 minutes.

Please replace the paragraphs at page 40, line 37 to page 41, line 6 with the following amended paragraph:

These peptides are:

Specific peptide of ULIP-1: GSARGSPTRPN (SEQ ID No. 37) (11 amino acids)

Specific peptide of ULIP-2: SSAKTSPAKQQA (SEQ ID No. 38) (12 amino acids)

Specific peptide of ULIP-3: PSAKSSPSKHQ (SEQ ID No. 39) (11 amino acids)

Specific peptide of ULIP-4: PARASCPGKIS (SEQ ID No. 40) (11 amino acids).

Please replace the Sequence Listing found in the specification on pages 45-59 (previously amended on October 9, 2003) with the substitute Sequence Listing provided herewith. A Statement Under 37 CFR § 1.821 and computer readable form of the sequence listing is also provided herewith.